PATENT

AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listing, of claims in the application.

Listing of the Claims

Claim 1 (currently amended): A method for differentiating DN \ species originating from cells of different individuals in a biological sample, the method comprising the steps of:

- (a) obtaining from one of the individuals a biological sampl; comprising the cells of different individuals, wherein the cells comprise the Dl A species; and
- (b) determining epigenetic differences detecting a methylati n difference between these the DNA species.

wherein detection of a methylation difference indicates DNA sp cies from different individuals.

Claim 2 (currently amended): A method according to claim 1 w terein the epigenetic difference is a difference in DNA methylation biological sample is a fluid or cellular sample or a mixture thereof.

Claim 3 (original): A method according to claim 1 wherein the l iological sample is plasma or serum.

Claim 4 (original): A method according to claim 1 wherein the t ological sample is blood.

PATENT

Claim 5 (original): A method according to claim 1 wherein on · of the individuals is a pregnant female and the other individual is an unborn fetus.

Claim 6 (original): A method according to claim 1 wherein on of the individuals is a transplantation recipient and the other individual is an organ d nor.

Claim 7 (original): A method according to claim 6 wherein the transplantation is a bone marrow transplantation.

Claim 8 (currently amended): A method according to claim 1 f in ther comprising the step of:

(c) measuring emecatrations the concentration of the DNA species having an epigenetic difference.

Claim 9 (canceled)

Claim 10 (currently amended): A method according to claim 2 further comprising the step of:

(c) adding sodium bisulfite to the biological sample or to the DNA species to detect a DNA methylation difference.

Claim 11 (currently amended): A method according to claim 2 further comprising the step of:

(c) performing a methylation-specific polymerase chain reaction to detect a DNA methylation difference.

Claim 12 (currently amended): A method according to claim 10 1 further comprising the steps of:

PATENT

- (c) amplifying the DNA species to generate a PCR produc and
- (d) sequencing DNA to detect a DNA methylation differen e the <u>PCR</u> product.

Claim 13 (currently amended): A method according to claim 4) I further comprising the step of:

(c) performing primer extension to detect a DNA methylati m difference.

Claim 14 (original): A method according to claim 5 wherein the biological sample is maternal plasma or serum.

Claim 15 (currently amended): A method according to claim 1. further comprising the step of:

(c) measuring the concentration of fetal DNA in maternal p asma or serum.

Claim16 (original): A method according to claim 15 wherein the concentration of fetal DNA measured is used to predict, monitor or diagnose or prognosticate a disorder.

Claim 17 (currently amended): A method according to claim 5 /herein an epigenetic mark the methylation difference is associated with a fetal or maternal disorder.

Claim 18 (original): A method according to claim 17 wherein tl e disorder is a chromosomal aneuploidy.

<u>PATENT</u>

Claim 19 (original): A method according to claim 18 wherein he chromosomal aneuploidy is trisomy 21 (Down syndrome).

Claim 20 (original): A method according to claim 17 wherein he disorder is preeclampsia.

Claim 21 (original): A method according to claim 17 wherein the disorder is an imprinting disorder.

Claim 22 (currently amended): A method according to claim 2 17 wherein the disorder is Prader-Willi syndrome.

Claim 23 (currently amended): A method according to claim 2 17 wherein the disorder is Angelman syndrome.

Claim 24 (currently amended): A method according to claim 1. wherein an epigenetic the methylation difference detected in fetal cells in the placenta i used as a fetus-specific marker in maternal plasma or serum.

Claim 25 (currently amended): A method according to claim 6 in ther comprising the step of:

(c) measuring the concentrations concentration of organ dor or and transplantation recipient DNA.

Claim 26 (currently amended): A method according to claim 25 wherein the measurements concentration of organ donor and transplantation recipient D <u>VA</u> are is used to predict the clinical progress of the transplantation recipient.

PATENT

Claim 27 (currently amended): A method according to claim 1 wherein one individual of the individuals is male and the other individual is female.

Claim 28 (currently amended): A method according to claim 27 wherein the epigenetic marker methylation difference is detected on an inactivated X c iromosome of the female individual.

Claim 29 (currently amended): A method according to claim 2 \(\) wherein a methylated DNA sequences sequence on the inactivated X chromosome are is used to detect DNA originating from the female individual.

Claim 30 (currently amended): A method according to claim 1 wherein the epigenetic differences are methylation difference is analyzed inside cells.

Claim 31 (currently amended): A method according to claim 3 wherein the epigenetic differences-are methylation difference is analyzed using in-situ methylation-specific polymerase chain reaction.

Claim 32 (currently amended): A method according to claim 1 wherein the epigenetic differences are methylation difference is used to sort or isolate calls from the individuals.

Claim 33 (currently amended): A method according to claim 1 wherein the epigenetic differences are methylation difference is used to purify DNA from the individuals.

PATENT

Claim 34 (withdrawn): A kit for differentiating DNA species a riginating from different individuals in a biological sample comprising one or more reag :nts for ascertaining the methylation status of a species of DNA.

Claim 35 (withdrawn): A kit according to claim 34 wherein the reagent for ascertaining the methylation status of the maternal DNA is sodium bisulfit.

Claim 36 (withdrawn): A kit according to claim 34 further con prising one or more reagents for detecting the presence of DNA.

Claim 37 (withdrawn): A kit according to claim 34 further con prising one or more reagents for amplifying the amount of DNA present in the biological sample.

Claim 38 (withdrawn): A kit according to claim 34 further comprising one or more apparatuses for obtaining a DNA sample.

Claim 39 (new): A method according to claim 5 wherein the m thylation difference is detected in fetal cells in the placenta.